

## **Determining Distribution for Planning an Infusion**

### **RELATED APPLICATION DATA**

This application claims priority of U.S. Provisional Application No.  
5 60/437,368, filed on December 31, 2002, which is incorporated herein by  
reference in its entirety.

### **FIELD OF THE INVENTION**

The invention relates generally to the field of planning an infusion and,  
10 more particularly to a method for identifying advantageous and/or  
non-advantageous infusion regions as well as a method and a device for assisting  
planning for introducing an infusion fluid.

### **BACKGROUND OF THE INVENTION**

15 Various medical methods require therapeutic agents to be directly infused  
into the tissue, with the aim of achieving a broad and optimum homogeneity of the  
distribution of the infusion fluid in the tissue. While the administered agents are  
generally fluids, the term "infusion" can include the administering of, for example,  
any fluid or gaseous or solid substance or infusion agent, such as, for example,  
20 medicines, cells, genes, enzymes, proteins, antibodies, hormones, viruses or the  
like. These substances are generally introduced directly into a body or body  
tissue, for example, into a patient's brain. The substance can be supplied within a  
relatively short period of time, e.g., by injection, or over a longer period of time,  
e.g., at a continuous, or, as the case may be, variable supply rate of the  
25 substance.

A method and a device for the targeted release of a medicine using  
magnetic resonance image detection are known from U.S. Pat. No. 6,061,587.  
U.S. Pat No. 5,583,902 discloses a method and a device for predicting  
organ-specific contrast amplification in a patient before an injection. U.S. Pat. No.  
30 5,720,720 discloses a method for micro-infusing at high flow rates, enabling  
agents to be released into the brain and other fixed tissue structures with  
convection-amplification. U.S. Pat. No. 5,205,289 describes an optimized dosage

administering system using graphic simulation techniques and computer-assisted, numerical optimization. U.S. Pat. No. 3,690,318 discloses a fluid infusion device comprising variable flow regulating means. A method and a device for nuclear spin flow image detection are known from U.S. Pat. No. 5,195,524. U.S. Pat. No. 5,840,026 describes a contrast medium supply system, which starts setting the contrast medium concentration and the injection parameters before or during an injection.

The homogeneity of the distribution of an infusion or of an infusion fluid can deteriorate if the infusion agent is introduced into a region in which the agent is transported through directional channels, which are not in themselves the infusion target, nor their end points. Instead of diffusing into the actual target areas, the infusion agent runs off along these "tracks", without achieving the desired effect.

## **SUMMARY OF THE INVENTION**

It is an object of the present invention to achieve increased control over the distribution of the infusion agent. One object of the invention is to prevent the infusion insertion point from coming too near to directional channels in which the infusion agent is quickly transported away and thus runs off without any effect.

According to one aspect of the invention, the invention is directed to a method for identifying advantageous and/or non-advantageous infusion regions in a tissue. Functional and/or structural anatomical data can be captured and the anatomical data can be evaluated with computer assistance. The anatomical data can be evaluated with respect to the distribution information contained therein, such as directional and/or velocity information. This can be used, for example, to identify, before the infusion fluid is introduced, the directional channels at which infusion fluid may be expected to be rapidly transported away when it is introduced. The basis for this is formed by anatomical data such as can be determined, for example, by an imaging system, such as a nuclear spin tomograph, a computer tomograph or similar known imaging systems. In this way, it is possible both to capture structural anatomical data, i.e., only data on the

tissue structure, and to obtain functional information, for example, data on certain regions having a specific function (auditory cortex, visual cortex, etc.) in the brain.

Using this information, it is then possible with computer assistance to determine which regions of the tissue contain transport pathways. In this way, it is also possible to find out whether the infusion agent will run off along so-called "tracks" without any effect if it is introduced at a particular point, or whether there will be a homogeneous diffusion into the surrounding tissue. It is therefore possible in this way, even before performing an infusion, to distinguish target areas having advantageous distribution properties from those having less advantageous or non-advantageous distribution properties.

The structural and/or anatomical properties of the tissue can change during the infusion. Such change can result from the physical and/or biochemical conditions of the infusion itself or from a reaction between the infused material and the tissue. In order to take these changes into account in their chronological course, corresponding adjustments can be made to the distribution information and made available to the user.

In one embodiment, the method for identifying transport pathways is based on diffusion measurements. These measurements can be modified in a mathematical algorithm by forming mean values of interference signals and converted into direction-independent (modified) or direction-dependent velocity information per volume element. Representations of the isotropy and/or anisotropy can likewise be based on diffusion measurements and can contain information on the direction-dependence of the fluid-transparency (permeability) of a volume element. The velocity information and isotropy information can be further used, each alone or combined with each other. Furthermore, the velocity information and/or isotropy information can be combined with other anatomical data, to increase or specify its significance.

In one embodiment, the velocity of diffusion of a volume element in the tissue can be determined within the framework of evaluating the distribution information, such as, for example, by identifying regions having rapid diffusion.

The distribution information, such as the diffusion velocity and isotropy, can be determined two-dimensionally, i.e., on the basis of two-dimensional image

information. If a number of such two-dimensional image information data sets are available on the anatomical structure, which in their planes make information on the distribution of the infusion fluid accessible, then these two-dimensional data sets can be combined to obtain three-dimensional distribution information.

5 Alternatively, there exists the possibility of directly determining three-dimensional image data sets and evaluating them with respect to their distribution information.

In accordance with another aspect, the invention relates to a method for assisting planning for introducing an infusion fluid into regions of the brain, wherein infusion regions are identified using a method such as been described  
10 above. Furthermore, introducing the infusion at a selected point is also planned and/or carried out by means of medical, e.g., stereotactic navigation. The person positioning the infusion can then be shown, with the aid of medical navigation, where the target of the infusion device is supposed to be. While positioning the infusion he can be guided in shifting the same until the optimum introduction  
15 target is reached. The infusion instruments can be tracked by a camera system, or magnetically by known tracking methods, and their spatial location can be shown in relation to the patient's anatomy on an image output.

It is possible within the framework of the aforementioned method to combine anatomical, functional and/or structural tissue data with information on  
20 the distribution of the infusion fluid to be expected. In other words, the anatomical patient data determined for simulating the distribution can also be used during navigation by being referenced and/or registered in the navigation system.

In accordance with another aspect, the present invention relates to a device for assisting planning for introducing an infusion fluid into regions of the  
25 brain. The device can include an imaging device, such as a nuclear spin tomograph, for capturing functional and/or structural anatomical data. A computer which, on the basis of the captured anatomical data, can produce an evaluation of the distribution information of an infusion fluid when it is introduced at particular points. The device can include computer-assisted, medical planning  
30 and navigation system for assisting in positioning an infusion device. Evaluating the distribution information and navigating can be assisted by a single computer system or by separate computer systems. Such a device enables the present

invention to be performed and adapted, with the aforementioned advantages with respect to the distribution of the infusion agent.

The imaging device, the computer or computers and the navigation system can be connected to each other via data connections, for a constant or retrievable exchange of data. It is possible to connect individual devices or all of these devices to each other in this way.

Furthermore, the invention also relates to a program which, when run on a computer or loaded onto a computer, causes the computer to perform a method as described above, and to a computer program storage medium comprising such a program.

### **BRIEF DESCRIPTION OF DRAWINGS**

These and further features of the present invention will be apparent with reference to the following description and drawings, wherein:

Figure 1 is a diagrammatic illustration of a system for identifying advantageous and non-advantageous infusion regions and/or for assisting planning for introducing a fluid in accordance with the present invention.

### **DETAILED DESCRIPTION OF THE INVENTION**

It is to be appreciated that the term "infusion", as used herein is intended to include the administering of, for example, any fluid or gaseous or solid substance or infusion agent, such as for example medicines, cells, genes, enzymes, proteins, antibodies, hormones, viruses or the like.

With reference to Figure 1, a device or system for identifying advantageous and non-advantageous infusion regions and/or for assisting planning for introducing a fluid is provided. The system includes an imaging device 1, such as a nuclear spin tomograph with a patient couch. A patient can be introduced into the nuclear spin tomograph, for example, via the head, and structural and/or functional anatomical data can be captured. The captured structural and/or functional anatomical data can be used to determine the structure of the head, for example, the brain. Once the recorded images have been processed, which can simply be performed in a processor in the nuclear spin tomograph, information is

then available on the structure of the patient's brain. In one embodiment, this data is recorded in such a way that brain structure can be registered and/or referenced in a medical navigation system, such as is indicated by the reference numeral 3. The medical navigation system can include an optical navigation  
5 system, such as is described in co-owned U.S. Pat. No. 6,351,659, which is incorporated herein by reference in its entirety.

In this embodiment, before the nuclear spin tomograph is recorded, markings can be attached to the patient's head. These markings can be identified both in the nuclear spin images and when tracked using the navigation system,  
10 such that the positional relationship between the markings and the detected regions of the brain structure is established and can also be used later in the course of the navigated positioning of the infusion.

A data connection 13 can be provided for exchanging data between the nuclear spin tomograph 1 and the navigation system 3. The data connection can  
15 include, for example, a data line, radio data transmission, or data transmission via transfer by storage media. The connection can thus be an on-line (continuous) connection, but can also be a retrievable (off-line) connection.

A processor or computer system 2 can exchange data with the nuclear spin tomograph 1 via an on-line or off-line connection 12. The functional and/or  
20 structural anatomical data from the nuclear spin tomograph 1, determined as described above, can be communicated to the computer 2 via the line 12. A program can then start in the computer, which on the basis of said anatomical data, evaluates and/or displays the distribution information and/or simulates the distribution of an infusion fluid at particular points. This will reveal that at some  
25 points, at which directional and/or rapid channels (tracks) are present, there can be a non-advantageous distribution, i.e., rapid run-off into undesired areas, if these are selected as the infusion point. On the other hand, target points for the infusion can be identified, at which a homogenous distribution of the infusion fluid may be expected. The computer 2 can make the advantageous points  
30 distinguishable from the non-advantageous ones, for example, by inserting corresponding indicators in the image data sets produced.

Using this information on advantageous and non-advantageous infusion regions in the brain tissue, one or more positions for the infusion device can then be planned by the user, such that a maximally homogenous distribution of the infusion fluid may be expected. To this end, the user of the computer 2 inputs the desired positions for the infusion device with the aid of a user interface and taking into account the information presented to him on the computer 2. With the aid of the distribution information contained in the computer, advantageous target points for the positions of the infusion device can also be suggested by the computer. Once this procedure has been concluded, the computer 2 transfers the positions for the infusion devices and/or other information to the navigation system 3 via an interface 23.

In addition to the navigation system described and incorporated above, the navigation system can be a known optical navigation system, such as is, for example, described in DE 196 396 152, the disclosure of which is incorporated herein its entirety by reference. Such a navigation system 3, which positionally tracks and registers the patient and medical instruments, for example, the infusion device, and displays them on an image output, referenced to determined image data, is shown in the drawing as an optical, camera-assisted system. Other navigation systems, including but not limited to magnetic or inductive navigation systems based on tracking magnetic and/or inductive signal emitters in the magnetic and/or electrical fields may be used.

When positioning the infusion, the navigation system 3 is placed near the patient or the patient is moved to it. The navigation system 3, which benefits from the anatomical data of the patient communicated via the line 12 and/or 23, can spatially assign this anatomy. This is due to the fact that the patient is wearing corresponding markings and/or natural landmarks, which have been both positionally recorded by the nuclear spin tomograph 1 and can be positionally detected by the navigation system using its camera system. Furthermore, the navigation system 3 can positionally track an infusion device provided with corresponding markings, and incorporate it such that it is clear to the person carrying out the treatment where the tip of the infusion device is currently situated with respect to the patient's anatomy. The navigation system 3 obtains the third

piece of information required, i.e., the data on advantageous and/or non-advantageous infusion regions in the tissue, from the computer 2 via the data line 23. With the aid of this data, specific regions in the brain, which are or are not advantageous for the infusion insertion point can be distinguished. When  
5 positioning the infusion, it can then be shown on a two-dimensional or three-dimensional view on the image output of the navigation system 2 where the tip of the infusion device is currently situated and whether this point promises a homogeneous distribution of the infusion fluid or not. When a physician recognizes on the screen output that he has just reached an advantageous  
10 infusion region with the tip of his instrument, the instrument can be placed there.

Accordingly, a method and a device which prevent infusion openings for infusion devices from being placed at unsuitable points in the tissue for releasing the infusion fluid are provided. This provides increased control over the effectiveness of the distribution of the infusion agent. The agent can, for  
15 example, be highly effectively injected directly into particular tumors. Because the infusion agent is prevented from running off without any effect, it is possible to use smaller quantities of the infusion agent and, because of the enabled homogeneous distribution, to generate an optimum effect. Furthermore, other regions of the brain and/or the body as a whole are prevented from being  
20 damaged by infusion fluid running off in an uncontrolled manner.

Although particular embodiments of the invention have been described in detail, it is understood that the invention is not limited correspondingly in scope, but includes all changes, modifications and equivalents coming within the spirit and terms of the claims appended hereto.